

REMARKS

Claims 112-141, 151-168, 185-216 and 226-243 are currently pending in the application. The foregoing separate sheets marked as "Listing of Claims" show all the claims in the application, with an indication of the current status of each.

Applicant thanks Examiner for the indication in the Office Action dated 10/15/2004 that all claims in the application are allowed. Further to Examiner's comments in that Office Action, Applicant has hereby amended the specification by replacing the first paragraph of the specification with a replacement paragraph that recites that the parent application of the present application is now abandoned and is no longer copending.

Further, Applicant herewith submits a corrected Sequence Listing for the application in which the sequences on page 89 and the sequences on page 69 are listed as SEQ ID NOS: 15 and 16 and SEQ ID NOS: 17 and 18, respectively. In addition, the specification has been amended in order to replace the two paragraphs that contain the sequences, paragraph [0380] on page 89 and paragraph [0329] on page 69, with replacement paragraphs that contain the SEQ ID NOS. Applicant submits that these amendments to the specification do not introduce any new matter, and requests entry of the replacement paragraphs. Applicant notes that the original paragraph [0380] contained underlining in the text within the sequence (the first 23 nucleotides) to indicate the T7 promoter sequence. This underlining has been retained in order to remain true to the original text, and should not be mistaken for new text, even though the new SEQ ID NOS. are also underlined to show that they are being added.

In view of the foregoing, Applicant submits that the application is now in *prima facie* condition for allowance. Should the Examiner find the application to be other than in condition for allowance, the Examiner is requested to contact the undersigned at 703-787-9400 (fax: 703-787-7557; email: ruth@wcc-ip.com) to discuss any other changes deemed necessary in a telephonic or personal interview.

If an extension of time is required for this response to be considered as being timely filed, a conditional petition is hereby made for such extension of time. Please charge any deficiencies in fees and credit any overpayment of fees to Attorney's Deposit Account No. 50-2041.

Respectfully submitted,

A handwritten signature in black ink, appearing to read 'Ruth E. Tyler-Cross', with a large, stylized flourish extending from the end of the signature.

Ruth E. Tyler-Cross
Reg. No. 45,922

Whitham, Curtis & Christofferson, P.C.
11491 Sunset Hills Road, Suite 340
Reston, VA 20190
703-787-9400
703-787-7557 (fax)



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re patent application of Betenbaugh et al.

Serial number: 09/930,440

Group Art Unit: 1652

Attorney Docket Number: 03940077pa

Examiner: Rao

Filed: 2001-08-16

For: ***"ENGINEERING INTRACELLULAR SIALYLATION PATHWAYS"***

SUBMISSION OF SUBSTITUTE SEQUENCE LISTING AND STATEMENT TO SUPPORT
FILING IN ACCORDANCE WITH 37 C.F.R. § 1.821-1.825

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Dear Sir:

In response to an Office Action mailed 10/15/2004, we enclose herein a corrected substitute computer readable form (diskette) and a corrected substitute paper copy of the sequence listings for the above-identified patent application. Please replace the Sequence Listing of the application with this substitute Sequence Listing. Also enclosed is a verified statement that the content of the paper and computer readable copies are the same and include no new matter.

Respectfully submitted,

Ruth E. Tyler Cross
Registration No. 45,922

Whitham, Curtis & Christofferson
11491 Sunset Hills Road; Suite 340
Reston, VA 20190
703-787-9400



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re patent application of Betenbaugh et al.

Serial number: 09/930,440

Attorney Docket Number: 03940077pa

Filed: 2001-08-16

For: ***"ENGINEERING INTRACELLULAR SIALYLATION PATHWAYS"***

STATEMENT TO SUPPORT FILING AND SUBMISSION IN ACCORDANCE WITH
37 C.F.R. §§ 1.821-1.825

Assistant Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450
Mail Stop SEQUENCE

Dear Sir:

In connection with a Substitute Sequence Listing submitted concurrently herewith, the undersigned states that:

1. the submission, filed herewith in accordance with 37 C.F.R. § 1.821 (g), does not include new matter;
2. the content of the attached paper copy and the attached computer readable copy of the Sequence Listing, submitted in accordance with 37 C.F.R. § 1.821(c) and (e), respectively, are the same; and
3. all statements made herein of their own knowledge are true and that all statements made on information and belief are believed to be true; and further, that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that

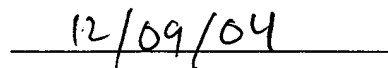
such willful false statements may jeopardize the validity of the application or any patent resulting therefrom.

Respectfully submitted,

A handwritten signature in black ink, appearing to read "Ruth E. Tyler-Cross", written over a horizontal line.

Ruth E. Tyler-Cross

Reg. No. 45,922

A handwritten date "12/09/04" in black ink, written over a horizontal line.

Date



Sequence Listing.ST25.txt
SEQUENCE LISTING

<110> Betenbaugh, Michael J.
Lawrence, Shawn J.
Lee, Yuan C.
Coleman, Timothy A.

<120> Engineering Intracellular Sialylation Pathways

<130> 03940077pa

<140> 09/930,440

<141> 2001-08-16

<150> US 60/122,582

<151> 1999-03-02

<150> US 60/169,624

<151> 1999-12-08

<150> US 60/227,579

<151> 2000-08-25

<150> US 09/516,793

<151> 2000-03-01

<160> 18

<170> PatentIn version 3.2

<210> 1

<211> 1429

<212> DNA

<213> Homo sapiens

<400> 1

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cagggagtga agaacatttt tgtgaatggc acaacaggag aaggcctgtc cctgagcgtc      180
tcagagcgtc gccaggttgc agaggagtgg gtgacaaaag ggaaggacaa gctggatcag      240
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aaagatatcc tgattaattt cctaaaggaa gtggctgtct cgcgccctgc cctgccattt      420
tattactatc acattcctgc cttgacaggg gtaaagattc gtgctgagga gttgttggat      480
gggattctgg ataagatccc caccttccaa gggctgaaat tcagtatac agatctctta      540
gacttcgggc aatgtgttga tcagaatcgc cagcaacagt ttgctttcct ttttgggggtg      600
gatgagcaac tgttgagtgc tctggtgatg ggagcaactg gagcagtggg cagttttgta      660
tccagagatt tatcaacttt gttgtcaaac taggttttgg agtgtcacag accaaagcca      720
tcatgactct ggtctctggg attccaatgg gccaccccg gcttccactg cagaaagcct      780
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Sequence Listing.ST25.txt

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tcattcattt cacagatttt tttgtggaga aatttctgtt tatatggatg aaatggaatc    1140
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tctggttcct aatcctatatt taaagttgtc taattttaaa ccactataat atgtcttcat    1260
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tttaataacca aactgtaaca tgtctcaact gtatacaact caaaatacac cagctcattt    1380
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<212> PRT
<213> Homo sapiens

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20          25          30

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```

Tyr Val Asp Tyr Leu Val Lys Glu Gln Gly Val Lys Asn Ile Phe Val
35          40          45

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```

Asn Gly Thr Thr Gly Glu Gly Leu Ser Leu Ser Val Ser Glu Arg Arg
50          55          60

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```

Gln Val Ala Glu Glu Trp Val Thr Lys Gly Lys Asp Lys Leu Asp Gln
65          70          75          80

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Val Ile Ile His Val Gly Ala Leu Ser Leu Lys Glu Ser Gln Glu Leu
85          90          95

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Ala Gln His Ala Ala Glu Ile Gly Ala Asp Gly Ile Ala Val Ile Ala
100          105          110

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Pro Phe Phe Leu Lys Pro Trp Thr Lys Asp Ile Leu Ile Asn Phe Leu
115          120          125

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Lys Glu Val Ala Ala Ala Ala Pro Ala Leu Pro Phe Tyr Tyr Tyr His
130          135          140

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Sequence Listing.ST25.txt

Ile Pro Ala Leu Thr Gly Val Lys Ile Arg Ala Glu Glu Leu Leu Asp
145 150 155 160

Gly Ile Leu Asp Lys Ile Pro Thr Phe Gln Gly Leu Lys Phe Ser Asp
165 170 175

Thr Asp Leu Leu Asp Phe Gly Gln Cys Val Asp Gln Asn Arg Gln Gln
180 185 190

Gln Phe Ala Phe Leu Phe Gly Val Asp Glu Gln Leu Leu Ser Ala Leu
195 200 205

Val Met Gly Ala Thr Gly Ala Val Gly Ser Phe Val Ser Arg Asp Leu
210 215 220

Ser Thr Leu Leu Ser Asn Val Leu Glu Cys His Arg Pro Lys Pro Ser
225 230 235 240

Leu Trp Ser Leu Gly Phe Gln Trp Ala His Pro Gly Phe His Cys Arg
245 250 255

Lys Pro Pro Gly Ser Leu Leu Ile Val Leu Lys Leu Asn Arg Ala Trp
260 265 270

Ile Ser Phe Leu Ser Leu Ile Arg Met Glu Thr Trp Lys Leu Val Ala
275 280 285

Ser Ala Ser Leu Ser Asn Gln Gly Phe Ala Pro Leu Arg His Asn Leu
290 295 300

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<211> 1305
<212> DNA
<213> Homo sapiens

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aagaacatta agcacctggc gggggtccc ctcattggct gggtcctgcg tgcggccctg 240
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gccaaacaat ttggtgcaca agttcatcga agaagttctg aagtttcaaa agacagctct 360
acctcactag atgcatcat agaatttctt aattatyata atgaggktga cattgtagga 420
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Sequence Listing.ST25.txt

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cggcctcgtc gacaagactg ggatggagaa ttatatgaaa atggctcatt ttattttgct 660
aaaagacatt tgatagagat gggttacttg cagggtgga aaatggcata ctacgaaatg 720
cgagctgaac atagtgtgga tatagatgtg gatattgatt ggcctattgc agagcaaaga 780
gtattaagat atggctatatt tggcaaagag aagcttaagg aaataaaaact tttggtttgc 840
aatattgatg gatgtctcac caatggccac atttatgtat caggagacca aaaagaaata 900
atatcttatg atgtaaaaga tgctattggg ataagtttat taaagaaaag tggatttgag 960
gtgaggctaa tctcagaaag ggcctgttca aagcagacgc tgtcttcttt aaaactggat 1020
tgcaaatgg aagtcagtgt atcagacaag ctagcagttg tagatgaatg gagaaaagaa 1080
atgggcctgt gctggaaaga agtggcatat cttggaaatg aagtgtctga tgaagagtgc 1140
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gttgataca tttgcaaatg taatgggtggc cgtggtgcca tccgagaatt tgcagagcac 1260
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<213> Homo sapiens

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<223> Xaa can be any naturally occurring amino acid

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1          5          10          15

```

```

Gly Arg Pro Ser Arg Gly Arg Pro Pro Lys Leu Gln Arg Asn Ser Arg
Page 4

```

Sequence Listing.ST25.txt

20

25

30

Gly Gly Gln Gly Arg Gly Val Glu Lys Pro Pro His Leu Ala Ala Leu
 35 40 45
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 50 55 60
 His Leu Ala Gly Val Pro Leu Ile Gly Trp Val Leu Arg Ala Ala Leu
 65 70 75 80
 Asp Ser Gly Ala Phe Gln Ser Val Trp Val Ser Thr Asp His Asp Glu
 85 90 95
 Ile Glu Asn Val Ala Lys Gln Phe Gly Ala Gln Val His Arg Arg Ser
 100 105 110
 Ser Glu Val Ser Lys Asp Ser Ser Thr Ser Leu Asp Ala Ile Ile Glu
 115 120 125
 Phe Leu Asn Tyr Xaa Asn Glu Xaa Asp Ile Val Gly Asn Ile Gln Ala
 130 135 140
 Thr Ser Xaa Cys Leu His Pro Thr Asp Leu Gln Lys Val Ala Glu Met
 145 150 155 160
 Ile Arg Glu Glu Gly Tyr Asp Ser Xaa Phe Ser Val Val Arg Arg His
 165 170 175
 Gln Phe Arg Trp Ser Glu Ile Gln Lys Gly Val Arg Glu Val Thr Glu
 180 185 190
 Pro Leu Asn Leu Asn Pro Ala Lys Arg Pro Arg Arg Gln Asp Trp Asp
 195 200 205
 Gly Glu Leu Tyr Glu Asn Gly Ser Phe Tyr Phe Ala Lys Arg His Leu
 210 215 220
 Ile Glu Met Gly Tyr Leu Gln Gly Gly Lys Met Ala Tyr Tyr Glu Met
 225 230 235 240
 Arg Ala Glu His Ser Val Asp Ile Asp Val Asp Ile Asp Trp Pro Ile
 245 250 255
 Ala Glu Gln Arg Val Leu Arg Tyr Gly Tyr Phe Gly Lys Glu Lys Leu
 260 265 270

Sequence Listing.ST25.txt

Lys Glu Ile Lys Leu Leu Val Cys Asn Ile Asp Gly Cys Leu Thr Asn
275 280 285

Gly His Ile Tyr Val Ser Gly Asp Gln Lys Glu Ile Ile Ser Tyr Asp
290 295 300

Val Lys Asp Ala Ile Gly Ile Ser Leu Leu Lys Lys Ser Gly Ile Glu
305 310 315 320

Val Arg Leu Ile Ser Glu Arg Ala Cys Ser Lys Gln Thr Leu Ser Ser
325 330 335

Leu Lys Leu Asp Cys Lys Met Glu Val Ser Val Ser Asp Lys Leu Ala
340 345 350

Val Val Asp Glu Trp Arg Lys Glu Met Gly Leu Cys Trp Lys Glu Val
355 360 365

Ala Tyr Leu Gly Asn Glu Val Ser Asp Glu Glu Cys Leu Lys Arg Val
370 375 380

Gly Leu Ser Gly Ala Pro Ala Asp Ala Cys Ser Tyr Ala Gln Lys Ala
385 390 395 400

Val Gly Tyr Ile Cys Lys Cys Asn Gly Gly Arg Gly Ala Ile Arg Glu
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Gln Lys

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cgcatggcca aggagtgtgg ggctgattgt gccaagttcc agaagagtga gctagaattc 180
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tacgccgagg aggttgggat cttcttcact gcctctggca tggatgagat ggcagttgaa 360
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Sequence Listing.ST25.txt

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gcctatcctc ctgaagacat ctttaatcta gtgggcaaga aggtcctggg cactgttgaa 1020
gaggatgaca ccatcatgga agaattggtg gataatcatg gcaaaaaaat caagtcttaa 1080
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<210> 6
<211> 359
<212> PRT
<213> Homo sapiens
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<400> 6
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```

```
His Pro Cys Phe Ile Ile Ala Glu Ile Gly Gln Asn His Gln Gly Asp
20 25 30
```

```
Leu Asp Val Ala Lys Arg Met Ile Arg Met Ala Lys Glu Cys Gly Ala
35 40 45
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```
Asp Cys Ala Lys Phe Gln Lys Ser Glu Leu Glu Phe Lys Phe Asn Arg
50 55 60
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```
Lys Ala Leu Glu Arg Pro Tyr Thr Ser Lys His Ser Trp Gly Lys Thr
65 70 75 80
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```
Tyr Gly Glu His Lys Arg His Leu Glu Phe Ser His Asp Gln Tyr Arg
85 90 95
```

```
Glu Leu Gln Arg Tyr Ala Glu Glu Val Gly Ile Phe Phe Thr Ala Ser
100 105 110
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Gly Met Asp Glu Met Ala Val Glu Phe Leu His Glu Leu Asn Val Pro
115 120 125
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Sequence Listing.ST25.txt

Phe Phe Lys Val Gly Ser Gly Asp Thr Asn Asn Phe Pro Tyr Leu Glu
130 135 140

Lys Thr Ala Lys Lys Gly Arg Pro Met Val Ile Ser Ser Gly Met Gln
145 150 155 160

Ser Met Asp Thr Met Lys Gln Val Tyr Gln Ile Val Lys Pro Leu Asn
165 170 175

Pro Asn Phe Cys Phe Leu Gln Cys Thr Ser Ala Tyr Pro Leu Gln Pro
180 185 190

Glu Asp Val Asn Leu Arg Val Ile Ser Glu Tyr Gln Lys Leu Phe Pro
195 200 205

Asp Ile Pro Ile Gly Tyr Ser Gly His Glu Thr Gly Ile Ala Ile Ser
210 215 220

Val Ala Ala Val Ala Leu Gly Ala Lys Val Leu Glu Arg His Ile Thr
225 230 235 240

Leu Asp Lys Thr Trp Lys Gly Ser Asp His Ser Ala Ser Leu Glu Pro
245 250 255

Gly Glu Leu Ala Glu Leu Val Arg Ser Val Arg Leu Val Glu Arg Ala
260 265 270

Leu Gly Ser Pro Thr Lys Gln Leu Leu Pro Cys Glu Met Ala Cys Asn
275 280 285

Glu Lys Leu Gly Lys Ser Val Val Ala Lys Val Lys Ile Pro Glu Gly
290 295 300

Thr Ile Leu Thr Met Asp Met Leu Thr Val Lys Val Gly Glu Pro Lys
305 310 315 320

Ala Tyr Pro Pro Glu Asp Ile Phe Asn Leu Val Gly Lys Lys Val Leu
325 330 335

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<212> DNA

Sequence Listing.ST25.txt

<213> Escherichia coli

<400> 7

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gaaaaaata taacaacaaa aagacctggt aatggtatca gtccgatgga gtggtataat    960
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<211> 346

<212> PRT

<213> Escherichia coli

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1           5           10           15

Ser Val Asp Ile Ala Arg Glu Met Ile Leu Lys Ala Lys Glu Ala Gly
          20           25           30

Val Asn Ala Val Lys Phe Gln Thr Phe Lys Ala Asp Lys Leu Ile Ser
          35           40           45

Ala Ile Ala Pro Lys Ala Glu Tyr Gln Ile Lys Asn Thr Gly Glu Leu
          50           55           60

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Sequence Listing.ST25.txt

Glu Ser Gln Leu Glu Met Thr Lys Lys Leu Glu Met Lys Tyr Asp Asp
 65 70 75 80
 Tyr Leu His Leu Met Glu Tyr Ala Val Ser Leu Asn Leu Asp Val Phe
 85 90 95
 Ser Thr Pro Phe Asp Glu Asp Ser Ile Asp Phe Leu Ala Ser Leu Lys
 100 105 110
 Gln Lys Ile Trp Lys Ile Pro Ser Gly Glu Leu Leu Asn Leu Pro Tyr
 115 120 125
 Leu Glu Lys Ile Ala Lys Leu Pro Ile Pro Asp Lys Lys Ile Ile Ile
 130 135 140
 Ser Thr Gly Met Ala Thr Ile Asp Glu Ile Lys Gln Ser Val Ser Ile
 145 150 155 160
 Phe Ile Asn Asn Lys Val Pro Val Gly Asn Ile Thr Ile Leu His Cys
 165 170 175
 Asn Thr Glu Tyr Pro Thr Pro Phe Glu Asp Val Asn Leu Asn Ala Ile
 180 185 190
 Asn Asp Leu Lys Lys His Phe Pro Lys Asn Asn Ile Gly Phe Ser Asp
 195 200 205
 His Ser Ser Gly Phe Tyr Ala Ala Ile Ala Ala Val Pro Tyr Gly Ile
 210 215 220
 Thr Phe Ile Glu Lys His Phe Thr Leu Asp Lys Ser Met Ser Gly Pro
 225 230 235 240
 Asp His Leu Ala Ser Ile Glu Pro Asp Glu Leu Lys His Leu Cys Ile
 245 250 255
 Gly Val Arg Cys Val Glu Lys Ser Leu Gly Ser Asn Ser Lys Val Val
 260 265 270
 Thr Ala Ser Glu Arg Lys Asn Lys Ile Val Ala Arg Lys Ser Ile Ile
 275 280 285
 Ala Lys Thr Glu Ile Lys Lys Gly Glu Val Phe Ser Glu Lys Asn Ile
 290 295 300
 Thr Thr Lys Arg Pro Gly Asn Gly Ile Ser Pro Met Glu Trp Tyr Asn
 305 310 315 320

Sequence Listing.ST25.txt

Leu Leu Gly Lys Ile Ala Glu Gln Asp Phe Ile Pro Asp Glu Leu Ile
 325 330 335

Ile His Ser Glu Phe Lys Asn Gln Gly Glu
 340 345

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 <213> Artificial
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 C,A,C/T,T,G,G,C,A,C/T,A/T/C,T,I,G,T,I,G,A

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 <223> n = t or c

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 <223> n = i

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 <223> n = c or t

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 <222> (12)..(12)
 <223> n = c or t

<220>
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 <222> (13)..(13)
 <223> n = a, c or t

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<220>
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 <222> (18)..(18)
 <223> n = i

<400> 9
 ntncantggc anntngtnga

20

<210> 10
 <211> 20
 <212> DNA
 <213> Artificial

Sequence Listing.ST25.txt

<220>
 <223> synthetic oligonucleotide primer
 G,A,G/A,A/T,T,A/C/T,G,A,C/T,I,I,I,C,C,I,G,G/C,I,C,A

<220>
 <221> misc_feature
 <222> (3)..(3)
 <223> n = g or a

<220>
 <221> misc_feature
 <222> (4)..(4)
 <223> n = a or t

<220>
 <221> misc_feature
 <222> (6)..(6)
 <223> n = a,c or t

<220>
 <221> misc_feature
 <222> (9)..(9)
 <223> n = c or t

<220>
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 <222> (10)..(10)
 <223> n = i

<220>
 <221> misc_feature
 <222> (11)..(11)
 <223> n = i

<220>
 <221> misc_feature
 <222> (12)..(12)
 <223> n = i

<220>
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 <222> (15)..(15)
 <223> n is a, c, g, or t

<220>
 <221> misc_feature
 <222> (17)..(17)
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<220>
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 <222> (18)..(18)
 <223> n = i

<400> 10
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20

<210> 11
 <211> 20
 <212> DNA
 <213> Artificial

Sequence Listing.ST25.txt

<220>
 <223> synthetic oligonucleotide primer
 T,G,I,C/G,C,I,G,G,I,I,I,G/A,T,C,T/G/A,A,T/A,C/T,T,C

<220>
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 <223> n = i

<220>
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 <222> (4)..(4)
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<220>
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 <222> (9)..(9)
 <223> n = i

<220>
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 <222> (10)..(10)
 <223> n = i

<220>
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 <222> (11)..(11)
 <223> n = i

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 <222> (12)..(12)
 <223> n = g or a

<220>
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 <222> (15)..(25)
 <223> n = t, g or a

<220>
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 <222> (17)..(17)
 <223> n = t or a

<220>
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 <222> (18)..(18)
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 tgnnrngnn nntcnaantc

20

<210> 12
 <211> 20
 <212> DNA

Sequence Listing.ST25.txt

<213> Artificial

<220>

<223> synthetic oligonucleotide primer A, C/A/G, C/T,
T,C,G/A,T,C,I,C,C,I,C,C,I,I,I,G/A,T,G

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<221> misc_feature

<222> (2)..(2)

<223> n = c, a or g

<220>

<221> misc_feature

<222> (3)..(3)

<223> n = c or t

<220>

<221> misc_feature

<222> (6)..(6)

<223> n = g or a

<220>

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<222> (9)..(9)

<223> n = i

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<222> (12)..(12)

<223> n = i

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<222> (15)..(15)

<223> n = i

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<220>

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<222> (17)..(17)

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<220>

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<223> n = g or a

<400> 12

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20

<210> 13

<211> 54

<212> DNA

<213> Artificial

<220>

<223> synthetic oligonucleotide primer

Sequence Listing.ST25.txt

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<210> 14
<211> 34
<212> DNA
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<220>
<223> synthetic oligonucleotide primer

<400> 14
gtacggtacc ttattaagac ttgatttttt tgcc 34

<210> 15
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<212> DNA
<213> Artificial

<220>
<223> synthetic oligonucleotide primer

<400> 15
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<210> 16
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<212> DNA
<213> Artificial

<220>
<223> synthetic oligonucleotide primer

<400> 16
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<210> 17
<211> 14
<212> PRT
<213> Escherichia coli

<400> 17

Ile Ile Ala Ile Ile Pro Ala Arg Ser Gly Ser Lys Gly Leu
1 5 10

<210> 18
<211> 14
<212> PRT
<213> Homo sapiens

<400> 18

Leu Ala Ala Leu Ile Leu Ala Arg Gly Gly Ser Lys Gly Ile
1 5 10